

Form PTO-1449		U.S. Department of Commerce Patent and Trademark Office				Atty. Docket No. 48231-AZ-PCT-US JPW/AJM/MML		Serial No. Not Yet Known							
<b>INFORMATION DISCLOSURE CITATION</b> (Use several sheets if necessary)						Applicants: Greenwald and Levitan Filing Date _____ Group 1653									
<b>U.S. PATENT DOCUMENTS</b>															
Examiner Initial		Document Number							Date	Name	Class	Subclass	Filing Date if Appropriate		
		6	3	7	6	2	3	9	4/23/03	Baumeister					
		6	0	8	7	1	5	3	7/11/00	Greenwald					
		5	8	4	0	5	4	0	11/24/98	St. George-Hyslop					
<b>FOREIGN PATENT DOCUMENTS</b>															
		Document Number							Date	Country	Class	Subclass	Translation		
													Yes	No	
	WO	9	7	1	1	9	5	6	3/3/97	PCT					x
<b>OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)</b>															
		European Search Report, dated November 20, 2002;													
		PCT International Search Report, dated January 21, 1997;													
		PCT Written Opinion, dated July 15, 1997;													
		Bai C. et al. "SKP1 connects cell cycle regulators to the ubiquitin proteolysis machinery through a novel motif, the F-box," Cell 86:263-74 (1996);													
		Brenner S. "The genetics of Caenorhabditis elegans", Genetics. (1974) 77(1):71-94;													
		Daigle I. and Li C. "apl-1, a Caenorhabditis elegans gene encoding a protein related to the human beta-amyloid protein precursor" Proc. Natl. Acad. Sci. U.S.A. (1993) 90(24):12045-9;													
		Database dbEST, National Center for Biotechnology Information, National Library of Medicine, GenBank Accession No. H19012 (1995);													
		Database EMBL Accession No: U35660 (1995) Levitan D., Greenwald I., "Caenorhabditis elegans membrane protein (sel-12) mRNA" XP002176178;													
		Ellisen L.W. et al. "TAN-1, the human homolog of the Drosophila Notch gene, is broken by chromosomal translocations in T lymphoblastic neoplasms," Cell 66:649-61 (1991);													
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		Hedgecock E.M. and Herman R.K. "The ncl-1 gene and genetic mosaics of Caenorhabditis elegans" Genetics (1995) 141(3):989-1006;			
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		Hubbard J. et al. "Sel-10 negative regulator of lin-12 activity in C. elegans, encodes a member of the CDC4 family of proteins," Genes Dev. 11:3182-93 (1997);			
		Kimble J. "Alteration in cell lineage following laser ablation of cells in the somatic gonad of C. elegans," Dev. Biol. 87:286-300 (1981);			
		King R.W. et al. "How proteolysis drives the cell cycle," Science 274:1652-58 (1996);			
		Levitan D. and Greenwald I. "Facilitation of lin-12-mediated signalling by sel-12, a Caenorhabditis elegans S182 Alzheimer's disease gene" Nature (1995) 377(6547): 351-4;			
		Levy-Lahad E. et al. "Genomic structure and expression of STM2, the chromosome 1 familial Alzheimer disease gene" Genomics (1996) 34(2):198-204;			
		Levy-Lahad E. et al. "Candidate gene for the chromosome 1 familial Alzheimer's disease locus" Science (1995) 269(5226):973-7;			
		L'Hernault S.W. and Arduengo P.M. "Mutation of a putative sperm membrane protein in Caenorhabditis elegans prevents sperm differentiation but not its associated meiotic divisions" J. Cell. Biol. (1992) 119(1):55-68;			
		Li X. and Greenwald I. "HOP-1, a Caenorhabditis elegans presenilin, appears to be functionally redundant with SEL-12 presenilin and to facilitate LIN-12 and GLP-1 signaling" Proc. Natl. Acad. Sci. U.S.A. (1997) 94(22):12204-9;			
		Mello C.C. et al. "Efficient gene transfer in C.elegans: extrachromosomal maintenance and integration of transforming sequences" EMBO J. (1991) 10(12):3959-70;			
		Neer E.J. et al. "The ancient regulatory-protein family of WD-repeat proteins," Nature 371:297-300 (1994);			
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		Rogaev E. I. et al. "Familial Alzheimer's disease in kindreds with missense mutations in a gene on chromosome 1 related to the Alzheimer's disease type 3 gene" Nature (1995) 376(6543):775-8;					
		Seydoux G. and Greenwald I. "Cell autonomy of lin-12 function in a cell fate decision in C. elegans," Cell 57:1237-45 (1989);					
		Shen J. et al. "Skeletal and CNS defects in presenilin-1-deficient mice," Cell 89:629-39 (1997);					
		Sherrington R. et al. "Cloning of a gene bearing missense mutations in early-onset familial Alzheimer's disease" Nature (1995) 375(6534):754-60;					
		Stratagene Cloning Systems Catalog, 1993, pages 27, 31, 32, and 313;					
		Struhl G. et al., "Intrinsic activity of the Lin-12 and Notch intracellular domains in vivo" Cell (1993) 74(2):331-45;					
		Sundaram M. and Greenwald I. "Genetic and phenotypic studies of hypomorphic lin-12 mutants in Caenorhabditis elegans" Genetics (1993) 135(3):755-63;					
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		Wen C. et al. "spr-2, a suppressor of the egg-laying defect caused by loss of sel-12 presenilin in Caenorhabditis elegans, is a member of the SET protein subfamily" Proc. Natl. Acad. Sci. U.S.A. (2000) 97(26):14524-9;					
		Wilkinson H.A. and Greenwald I. "Spatial and temporal patterns of lin-12 expression during C. elegans hermaphrodite development" Genetics (1995) 141(2):513-26;					
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		Wilson R. et al. "2.2 Mb of contiguous nucleotide sequence from chromosome III of C. elegans," Nature 368:32-38 (1994);					
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Sequence Listing

Applicants submit herewith (a) a paper copy of the Sequence Listing, inserted into the specification following the Abstract of the Disclosure, (b) a request to use the computer readable form (CRF) of the sequence listing submitted in prior application U.S. Serial No. 09/043,944, attached hereto as **Exhibit A**, and (c) a statement in accordance with 37 C.F.R. §1.821(f) attached hereto as **Exhibit B**, certifying that the CRF and written sequence listing contain the same sequence information.

Information Disclosure

In accordance with their duty of disclosure under 37 C.F.R. §1.56, applicants request that the following disclosures be made of record in the above-identified application pursuant to 37 C.F.R. §1.97(b). These references are also listed on the Form PTO-1449 attached hereto as **Exhibit C**. Copies of these references were submitted in connection with prior application U.S. Serial No. 09/043,944.

1. U.S. Patent No. 6,376,239, issued April 23, 2002, Baumeister;
2. U.S. Patent No. 6,087,153, issued July 11, 2000, Greenwald et al.;
3. U.S. Patent No. 5,840,540, issued November 24, 1998, Peter H. St. George-Hyslop et al.;
4. PCT International Application No. WO 97/11956, published April 3, 1997;
5. European Search Report, dated November 20, 2002;
6. PCT International Search Report, dated January 21, 1997;
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8. Bai C. et al. "SKP1 connects cell cycle regulators to the ubiquitin proteolysis machinery through a novel motif, the F-box," *Cell* 86:263-74 (1996);
9. Brenner S. "The genetics of *Caenorhabditis elegans*", *Genetics*. (1974) 77(1):71-94;
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13. Ellisén L.W. et al. "TAN-1, the human homolog of the *Drosophila* Notch gene, is broken by chromosomal translocations in T lymphoblastic neoplasms," *Cell* 66:649-61 (1991);
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38. Stratagene Cloning Systems Catalog, 1993, pages 27, 31, 32, and 313;
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No fee, other than the enclosed filing fee, is deemed necessary in connection with the filing of this Preliminary Amendment. However, if any additional fee is required, authorization is hereby given to charge the amount of such fee to Deposit Account No. 03-3125.

If a telephone interview would be of assistance in advancing prosecution of the subject application, applicants' undersigned attorneys invite the Examiner to telephone them at the number provided below.

Respectfully submitted,

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John P. White  
Registration No. 28,678  
Alan J. Morrison  
Registration No. 37,399  
Attorneys for Applicants  
Cooper & Dunham, LLP  
1185 Avenue of the Americas  
New York, New York 10036  
(212)278-0400